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=> s betamethasone/cn
L1
             1 BETAMETHASONE/CN
=> d 11
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS on STN
L1
     378-44-9 REGISTRY
RN
     Pregna-1,4-diene-3,20-dione, 9-fluoro-11,17,21-trihydroxy-16-methyl-,
CN
     (11.beta., 16.beta.) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Pregna-1,4-diene-3,20-dione, 9-fluoro-11.beta.,17,21-trihydroxy-16.beta.-
     methyl- (8CI)
OTHER NAMES:
     .beta.-Methasone
CN
     .beta.-Methasone alcohol
     24: PN: US20030109453 SEQID: 23 claimed sequence
CN
CN
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     dione
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     diene-3,20-dione
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CN
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     Bedifos
CN
     Betacorlan
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     Betacortril
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     Betadexamethasone
     Betamethasone
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     Betamethazone
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     Betsolan
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     Bifas
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     Diprospan
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     Flubenisolone
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     Visubeta
FS
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MF
     C22 H29 F O5
CI
LC
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       BIOSIS, BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB,
       CHEMCATS, CHEMLIST, CIN, CSCHEM, DDFU, DIOGENES, DRUGU, EMBASE, HSDB*,
       IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, NIOSHTIC, PHAR,
       PHARMASEARCH, PIRA, PROMT, RTECS*, SPECINFO, TOXCENTER, USAN, USPAT2,
       USPATFULL, VETU
         (*File contains numerically searchable property data)
     Other Sources:
                      EINECS**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1823 REFERENCES IN FILE CA (1947 TO DATE)

43 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

1824 REFERENCES IN FILE CAPLUS (1947 TO DATE)

58 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

=> s exosurf/cn

L2 1 EXOSURF/CN

=> d 12

L2 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS on STN

RN 99732-49-7 REGISTRY

CN 3,5,9-Trioxa-4-phosphapentacosan-1-aminium, 4-hydroxy-N,N,N-trimethyl-10-oxo-7-[(1-oxohexadecyl)oxy]-, inner salt, 4-oxide, (7R)-, mixt. with formaldehyde polymer with oxirane and 4-(1,1,3,3-tetramethylbutyl)phenol and 1-hexadecanol (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 1-Hexadecanol, mixt. contg. (9CI)

CN 3,5,9-Trioxa-4-phosphapentacosan-1-aminium, 4-hydroxy-N,N,N-trimethyl-10-oxo-7-[(1-oxohexadecyl)oxy]-, inner salt, 4-oxide, (R)-, mixt. with formaldehyde polymer with oxirane and 4-(1,1,3,3-tetramethylbutyl)phenol and 1-hexadecanol

CN Formaldehyde, polymer with oxirane and 4-(1,1,3,3-tetramethylbutyl)phenol, mixt. contg. (9CI)

CN Oxirane, polymer with formaldehyde and 4-(1,1,3,3-tetramethylbutyl)phenol, mixt. contg. (9CI)

CN Phenol, 4-(1,1,3,3-tetramethylbutyl)-, polymer with formaldehyde and oxirane, mixt. contg. (9CI)

OTHER NAMES:

CN Exosurf

CN Surfexo

FS STEREOSEARCH

MF C40 H80 N O8 P . C16 H34 O . (C14 H22 O . C2 H4 O . C H2 O)x

CI MXS

PCT Phenolic resin, Polyether, Polyether formed

SR CA

LC STN Files: ADISNEWS, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CIN, DIOGENES, DRUGUPDATES, EMBASE, MEDLINE, MRCK*, PHAR, PROMT, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

CM 1

CRN 36653-82-4

 ${
m HO^-}$ (CH₂)₁₅ $^-{
m Me}$

CM 2

CRN 63-89-8 CMF C40 H80 N O8 P

Absolute stereochemistry. Rotation (+).

CM 3

CRN 25301-02-4

CMF (C14 H22 O . C2 H4 O . C H2 O) x

CCI PMS

CM 4

CRN 140-66-9 CMF C14 H22 O

CM 5

CRN 75-21-8 CMF C2 H4 O

 $^{\circ}$

CM 6

CRN 50-00-0 CMF C H2 O

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     108778-82-1 REGISTRY
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CN
OTHER NAMES:
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    A 60386X
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     Surfactant TA
     Surfacten
CN
CN
     Survanta
MF
     Unspecified
CI
     MAN
SR
     CA
                  ADISNEWS, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT,
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     STN Files:
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       MEDLINE, MRCK*, PHAR, PHARMASEARCH, PROMT, TOXCENTER, USAN, USPATFULL
         (*File contains numerically searchable property data)
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=> s BERACTANT/cn
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L4
=> d 14
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L4
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     108778-82-1 REGISTRY
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     Beractant (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
    A 60386X
CN
     Surfactant TA
CN
     Surfacten
CN
     Survanta
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     CA
LC
                  ADISNEWS, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT,
     STN Files:
       CAPLUS, CBNB, CIN, DIOGENES, DRUGNL, DRUGPAT, DRUGUPDATES, EMBASE, IPA,
       MEDLINE, MRCK*, PHAR, PHARMASEARCH, PROMT, TOXCENTER, USAN, USPATFULL
         (*File contains numerically searchable property data)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
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L5
=> s phosphatidylcholine/cn
             0 PHOSPHATIDYLCHOLINE/CN
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=> file medicine
FILE 'DRUGMONOG' ACCESS NOT AUTHORIZED
COST IN U.S. DOLLARS

SINCE FILE TOTAL SESSION 35.24 35.45

FULL ESTIMATED COST

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FILE 'SCISEARCH' ENTERED AT 13:09:07 ON 22 JUL 2003 COPYRIGHT 2003 THOMSON ISI

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FILE 'USPATFULL' ENTERED AT 13:09:07 ON 22 JUL 2003 CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 13:09:07 ON 22 JUL 2003
CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

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(FILE 'HOME' ENTERED AT 13:04:08 ON 22 JUL 2003)

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L2
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L4
              1 S BERACTANT/CN
L5
            603 S PHOSPHATIDYLCHOLINE
L6
              O S PHOSPHATIDYLCHOLINE/CN
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     CAPLUS, CEN, DGENE, DRUGB, DRUGLAUNCH, DRUGMONOG2, DRUGNL, DRUGU, EMBAL,
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           276 L7 AND L8
L9
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The previous command name entered was not recognized by the system.
For a list of commands available to you in the current file, enter
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L11
            11 L9 AND L10
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DUPLICATE IS NOT AVAILABLE IN 'ADISINSIGHT, ADISNEWS, DGENE, DRUGLAUNCH, DRUGMONOG2, KOSMET, MEDICONF, NUTRACEUT, PCTGEN, PHARMAML'. ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE PROCESSING COMPLETED FOR L11

11 DUP REM L11 (0 DUPLICATES REMOVED)

=> d 112 1-11 ibib, kwic

L12 ANSWER 1 OF 11 USPATFULL on STN

ACCESSION NUMBER:

2003:72168 USPATFULL

TITLE:

64 human secreted proteins

INVENTOR (S):

Ruben, Steven M., Olney, MD, UNITED STATES Rosen, Craig A., Laytonsville, MD, UNITED STATES Young, Paul E., Gaithersburg, MD, UNITED STATES Greene, John M., Gaithersburg, MD, UNITED STATES

Ni, Jian, Germantown, MD, UNITED STATES Feng, Ping, Gaithersburg, MD, UNITED STATES

Florence, Kimberly A., Rockville, MD, UNITED STATES Hu, Jing-Shan, Mountain View, CA, UNITED STATES Ferrie, Ann M., Tewksbury, MA, UNITED STATES Yu, Guo-Liang, Berkeley, CA, UNITED STATES Duan, Roxanne D., Bethesda, MD, UNITED STATES Janat, Fouad, Westerly, RI, UNITED STATES

NUMBER KIND DATE -----

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.: US 2003050455 A1 20030313 US 2001-776724 A1 20010206 (9)

Continuation-in-part of Ser. No. US 2000-669688, filed

on 26 Sep 2000, PENDING Continuation of Ser. No. US

1999-229982, filed on 14 Jan 1999, PENDING Continuation-in-part of Ser. No. WO 1998-US14613, filed

on 15 Jul 1998, UNKNOWN

| | | | NUMBER | DATE | |
|----------|--------------|----------------------------|--|--|--|
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| | | US | 1997-53441P | 19970722 | (60) |

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

Utility

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS:

23

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

2 Drawing Page(s)

LINE COUNT:

21934

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

. . . in certain tissues or cell types (e.g., pulmonary, DETD

developmental, reproductive, breast, and cancerous and wounded tissues)

or bodily fluids (e.g., pulmonary surfactant, lymph,

serum, plasma, urine, synovial fluid and spinal fluid) or another tissue

or cell sample taken from an individual having.

. . . a reservoir, such as an Ommaya reservoir. Pulmonary DETD administration can also be employed, e.g., by use of an inhaler or nebulizer, and formulation with an aerosolizing agent.

. . . Anti-inflammatory agents that may be administered with the Therapeutics of the invention include, but are not limited to, corticosteroids (e.g. betamethasone, budesonide, cortisone, dexamethasone, hydrocortisone, methylprednisolone, prednisolone, prednisone, and triamcinolone), nonsteroidal anti-inflammatory drugs (e.g., diclofenac, diflunisal, etodolac, fenoprofen, floctafenine, flurbiprofen, ibuprofen, . .

DETD

DETD

. . . (cosyntropin); adrenocortical steroids and their synthetic analogs such as ACLOVATE.TM. (alclometasone dipropionate), CYCLOCORT.TM. (amcinonide), BECLOVENT.TM. and VANCERIL.TM. (beclomethasone dipropionate), CELESTONE.TM. (betamethasone), BENISONE.TM. and UTICORT.TM. (betamethasone benzoate), DIPROSONE.TM. (betamethasone dipropionate), CELESTONE PHOSPHATE.TM. (betamethasone sodium phosphate), CELESTONE SOLUSPAN.TM. (betamethasone sodium phosphate and acetate), BETA-VAL.TM. and VALISONE.TM. (betamethasone valerate), TEMOVATE.TM. (clobetasol propionate), CLODERM.TM. (clocortolone pivalate), CORTEF.TM. and HYDROCORTONE.TM. (cortisol (hydrocortisone)), HYDROCORTONE ACETATE.TM. (cortisol (hydrocortisone) acetate), LOCOID.TM. (cortisol (hydrocortisone)).

L12 ANSWER 2 OF 11 USPATFULL on STN

ACCESSION NUMBER:

TITLE:

INVENTOR(S):

2003:71333 USPATFULL

186 human secreted proteins Ruben, Steven M., Olney, MD, UNITED STATES Rosen, Craig A., Laytonsville, MD, UNITED STATES Soppet, Daniel R., Centreville, VA, UNITED STATES Carter, Kenneth C., North Potomac, MD, UNITED STATES Bednarik, Daniel P., Columbia, MD, UNITED STATES Endress, Gregory A., Florence, MA, UNITED STATES Yu, Guo-Liang, Berkeley, CA, UNITED STATES Ni, Jian, Germantown, MD, UNITED STATES Feng, Ping, Gaithersburg, MD, UNITED STATES Young, Paul E., Gaithersburg, MD, UNITED STATES Greene, John M., Gaithersburg, MD, UNITED STATES Ferrie, Ann M., Painted Post, NY, UNITED STATES Duan, D. Roxanne, Bethesda, MD, UNITED STATES Hu, Jing-Shan, Mountain View, CA, UNITED STATES - Florence, Kimberly A., Rockville, MD, UNITED STATES Olsen, Henrik S., Gaithersburg, MD, UNITED STATES Fischer, Carrie L., Burke, VA, UNITED STATES Ebner, Reinhard, Gaithersburg, MD, UNITED STATES Brewer, Laurie A., St. Paul, MN, UNITED STATES Moore, Paul A., Germantown, MD, UNITED STATES Shi, Yanggu, Gaithersburg, MD, UNITED STATES LaFleur, David W., Washington, DC, UNITED STATES Li, Yi, Sunnyvale, CA, UNITED STATES Zeng, Zhizhen, Lansdale, PA, UNITED STATES Kyaw, Hla, Frederick, MD, UNITED STATES

| NUMBER | KIND | DATE |
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PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.: US 2003049618 A1 20030313 US 2001-809391 A1 20010316 (9) Continuation-in-part of Ser. No. US 1998-149476, filed on 8 Sep 1998, GRANTED, Pat. No. US 6420526 Continuation-in-part of Ser. No. WO 1998-US4493, filed

on 6 Mar 1998, UNKNOWN

NUMBER DATE

PRIORITY INFORMATION:

US 2000-190068P 20000317 (60) US 1997-40162P 19970307 (60)

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                 US 1997-56632P
                                    19970822 (60)
                 US 1997-56664P
                                    19970822 (60)
                 US 1997-56876P
                                    19970822 (60)
                 US 1997-56881P
                                    19970822 (60)
                 US 1997-56909P
                                    19970822 (60)
                 US 1997-56875P
                                    19970822 (60)
                 US 1997-56862P
                                    19970822 (60)
                 US 1997-56887P
                                    19970822 (60)
                                    19970822 (60)
                 US 1997-56908P
                 US 1997-48964P
                                    19970606 (60)
                                    19970905 (60)
                 US 1997-57650P
                 US 1997-56884P
                                    19970822 (60)
                 US 1997-57669P
                                    19970905 (60)
                 US 1997-49610P
                                    19970613 (60)
                 US 1997-61660P
                                    19971009 (60)
                 US 1997-51926P
                                    19970708 (60)
                 US 1997-52874P
                                    19970716 (60)
                 US 1997-58785P
                                    19970912 (60)
                 US 1997-55724P
                                    19970818 (60)
                 Utility
                 APPLICATION
                 HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,
                 ROCKVILLE, MD, 20850
                 23
                 1
                 2 Drawing Page(s)
                 26235
     . thymus, and other tissue of the immune system, and cancerous
and wounded tissues) or bodily fluids (e.g. lymph, amniotic fluid,
pulmonary surfactant, serum, plasma, urine, synovial
fluid or spinal fluid) or another tissue or cell sample taken from an
individual having such.
     . in certain tissues and cell types (e.g., lung, developing, and
cancerous and wounded tissues) or bodily fluids (e.g. amniotic fluid,
pulmonary surfactant, serum, plasma, urine, synovial
fluid or spinal fluid) or another tissue or cell sample taken from an
individual having such.
     . tissues and cell types (e.g. immune, blood cells and lung, and
cancerous and wounded tissues) or bodily fluids (e.g. lymph,
pulmonary surfactant, serum, plasma, urine, synovial
fluid or spinal fluid) or another tissue or cell sample taken from an
individual having such.
     . types (e.g., fetal tissue, pulmonary tissue, and melanocytes,
and cancerous and wounded tissues) or bodily fluids (e.g. lymph,
amniotic fluid, pulmonary surfactant, serum, plasma,
urine, synovial fluid or spinal fluid) or another tissue or cell sample
taken from an individual having such.
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a reservoir, such as an Ommaya reservoir. Pulmonary

administration can also be employed, e.g., by use of an inhaler or

19970905 (60)

US 1997-57761P

DOCUMENT TYPE:

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

LEGAL REPRESENTATIVE:

FILE SEGMENT:

LINE COUNT:

SITMM

SUMM

SUMM

SUMM

DETD

nebulizer, and formulation with an aerosolizing agent. DETD . . Anti-inflammatory agents that may be administered with the Therapeutics of the invention include, but are not limited to, corticosteroids (e.g. betamethasone, budesonide, cortisone, dexamethasone, hydrocortisone, methylprednisolone, prednisolone, prednisone, and triamcinolone), nonsteroidal anti-inflammatory drugs (e.g., diclofenac, diflunisal, etodolac, fenoprofen, floctafenine, flurbiprofen, ibuprofen, (cosyntropin); adrenocortical steroids and their synthetic DETD analogs such as ACLOVATE.TM. (alclometasone dipropionate), CYCLOCORT.TM. (amcinonide), BECLOVENT.TM. and VANCERIL.TM. (beclomethasone dipropionate), CELESTONE.TM. (betamethasone), BENISONE.TM. and UTICORT.TM. (betamethasone benzoate), DIPROSONE.TM. (betamethasone dipropionate), CELESTONE PHOSPHATE.TM. (betamethasone sodium phosphate), CELESTONE SOLUSPAN.TM. (betamethasone sodium phosphate and acetate), BETA-VAL.TM. and VALISONE.TM. (betamethasone valerate), TEMOVATE.TM. (clobetasol propionate), CLODERM.TM. (clocortolone pivalate), CORTEF.TM. and HYDROCORTONE.TM. (cortisol (hydrocortisone)), HYDROCORTONE ACETATE.TM. (cortisol (hydrocortisone) acetate), LOCOID.TM. (cortisol (hydrocortisone). L12 ANSWER 3 OF 11 USPATFULL on STN ACCESSION NUMBER: 2003:38352 USPATFULL TITLE: 143 human secreted proteins INVENTOR(S): Rosen, Craig A., Laytonsville, MD, UNITED STATES Ruben, Steven M., Olney, MD, UNITED STATES Moore, Paul A., Germantown, MD, UNITED STATES Young, Paul E., Gaithersburg, MD, UNITED STATES Komatsoulis, George A., Silver Spring, MD, UNITED Birse, Charles E., North Potomac, MD, UNITED STATES Duan, Roxanne D., Bethesda, MD, UNITED STATES Florence, Kimberly A., Rockville, MD, UNITED STATES Soppet, Daniel R., Centreville, VA, UNITED STATES KIND DATE NUMBER -----US 2003027999 A1 20030206 US 2001-986480 A1 20011108 PATENT INFORMATION: APPLICATION INFO.: (9) RELATED APPLN. INFO.: Continuation-in-part of Ser. No. WO 2000-US12788, filed on 11 May 2000, UNKNOWN NUMBER DATE -----US 1999-134068P 19990513 (60) PRIORITY INFORMATION: DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, ROCKVILLE, MD, 20850 NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: LINE COUNT: 29687 CAS INDEXING IS AVAILABLE FOR THIS PATENT. SUMM . . or cell types (e.g., neural, vascular, pulmonary, and cancerous and wounded tissues) or bodily fluids (e.g., lymph, serum, plasma, urine, pulmonary surfactant, sputum, lavage, synovial fluid and spinal fluid) or another tissue or cell sample taken from an individual having such a. SUMM . . . in certain tissues or cell types (e.g., pulmonary, muscle, and cancerous and wounded tissues) or bodily fluids (e.g., lymph, serum, pulmonary surfactant, sputum, plasma, urine, synovial fluid and spinal fluid) or another tissue or cell sample taken from an

individual having such.

```
. . . cell types (e.g., growth developmental, pulmonary, and
SUMM
       cancerous and wounded tissues) or bodily fluids (e.g., lymph, serum,
      plasma, amniotic fluid, pulmonary surfactant,
       pulmonary lavage, sputum, urine, synovial fluid and spinal fluid) or
       another tissue or cell sample taken from an individual having.
       . . . a reservoir, such as an Ommaya reservoir. Pulmonary
SUMM
      administration can also be employed, e.g., by use of an inhaler or
      nebulizer, and formulation with an aerosolizing agent.
            . Anti-inflammatory agents that may be administered with the
DETD
      Therapeutics of the invention include, but are not limited to,
       corticosteroids (e.g. betamethasone, budesonide, cortisone,
       dexamethasone, hydrocortisone, methylprednisolone, prednisolone,
      prednisone, and triamcinolone), nonsteroidal anti-inflammatory drugs
       (e.g., diclofenac, diflunisal, etodolac, fenoprofen, floctafenine,
       flurbiprofen, ibuprofen,. .
         . . (cosyntropin); adrenocortical steroids and their synthetic
DETD
       analogs such as ACLOVATE.TM. (alclometasone dipropionate), CYCLOCORT.TM.
       (amcinonide), BECLOVENT.TM. and VANCERIL.TM. (beclomethasone
       dipropionate), CELESTONE.TM. (betamethasone), BENISONE.TM. and
       UTICORT.TM. (betamethasone benzoate), DIPROSONE.TM. (
      betamethasone dipropionate), CELESTONE PHOSPHATE.TM. (
      betamethasone sodium phosphate), CELESTONE SOLUSPAN.TM. (
      betamethasone sodium phosphate and acetate), BETA-VAL.TM. and
       VALISONE.TM. (betamethasone valerate), TEMOVATE.TM.
       (clobetasol propionate), CLODERM.TM. (clocortolone pivalate), CORTEF.TM.
       and HYDROCORTONE.TM. (cortisol (hydrocortisone)), HYDROCORTONE
       ACETATE.TM. (cortisol (hydrocortisone) acetate), LOCOID.TM. (cortisol
       (hydrocortisone).
L12 ANSWER 4 OF 11 USPATFULL on STN
ACCESSION NUMBER:
                       2003:10272 USPATFULL
                       Pharmaceutical preparation for the inhalation of
TITLE:
                       antithrombin in inflammatory lung diseases and ARDS
                       Hoffmann, Johannes, Munchen, GERMANY, FEDERAL REPUBLIC
INVENTOR(S):
                       Wiedermann, Christian, Innsbruck, AUSTRIA
                       Roemisch, Juergen, Marburg, GERMANY, FEDERAL REPUBLIC
                           NUMBER
                                      KIND DATE
                       US 2003007966 A1 20030109
US 2002-188957 A1 20020705 (10)
PATENT INFORMATION:
APPLICATION INFO.:
                             NUMBER DATE
                       _______
PRIORITY INFORMATION: DE 2001-132307 20010706
DOCUMENT TYPE:
                       Utility
FILE SEGMENT:
                       APPLICATION
LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow,, Garrett & Dunner,
                       L.L.P., 1300 I Street, N.W., Washington, DC, 20005-3315
NUMBER OF CLAIMS:
EXEMPLARY CLAIM:
                       1
                       177
LINE COUNT:
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       . . . permeability represents an important part of the acute lung
       damage, both the chemical composition and the functional activity of the
      pulmonary surfactant being modified in patients with
      ARDS (2). These symptoms also occur in similar form in other
      inflammatory lung diseases.
       . . . compositions for the treatment of IRDS and ARDS have already
SUMM
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been specified which contain at least one glucocorticoid and one

mortality caused by these syndromes can be reduced by pharmaceuticals of

pulmonary surfactant. The treatment period and the

this type. From international.

DETD [0009] Also advantageous are pharmaceutical preparations which contain antithrombin III together with a pulmonary surfactant and/or with an antiinflammatory or a glucocorticoid selected from the group consisting of betamethasone, methylprednisolone and/or dexamethasone. The pulmonary surfactant is preferably a highly purified, natural surfactant made from homogenized porcine lungs or bovine lungs and phospholipids. Liquid pulmonary surfactant preparations are expediently lyophilized before or after the addition of the glucocorticosteroid and then micronized. Compositions according to the invention.

DETD [0012] In clinical investigations, the value of local nebulization of vasodilatory or antiinflammatory substances can be confirmed, where, for example, an improvement in the gas exchange short-term could be. . .

CLM What is claimed is:

. to 6, which contains antithrombin III together with pulmonary surfactants and/or with a glucocorticoid selected from the group consisting of **betamethasone**, methylprednisolone and/or dexamethasone.

TT 50-02-2, Dexamethasone 83-43-2, Methylprednisolone 378-44-9,
Betamethasone 9000-94-6, Antithrombin 9035-81-8, Antitrypsin
42617-41-4, Activated protein C 122320-05-2, Proteinase inhibitor, MPI
133249-66-8, Proteinase inhibitor, elafin 194554-71-7, Tissue factor
pathway inhibitor

(pharmaceutical prepn. for inhalation comprising antithrombin for treating inflammatory lung diseases and ARDS)

L12 ANSWER 5 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2003:148746 USPATFULL

TITLE: Composition and method for decreasing upper respiratory

airway resistance

INVENTOR(S): Mautone, Alan J., Morristown, NJ, United States

PATENT ASSIGNEE(S): Scientific Development and Research, Inc, Belleville,

NJ, United States (U.S. corporation)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-450884, filed

on 28 Nov 1999

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Krass, Frederick ASSISTANT EXAMINER: Jagoe, Donna

LEGAL REPRESENTATIVE: Strauss, Esq., Richard L.

NUMBER OF CLAIMS: 53 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT: 1336

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . other phospholipid, and the lysophospholipids; or any of the plasmalogens, dialklylphospholipids, phosphonolipids; carbohydrates and proteins, such as, for example, albumin, pulmonary surfactant proteins A, B, C and D. The naturally occurring surfactant system is further described in U.S. Pat. No. 5,306,483.

SUMM . . . to, glucose, fructose, galactose, pneumogalactan, dextrose, 0.5 to 10% by weight; and proteins such as, but not limited to albumin, pulmonary surfactant specific proteins A, B, C, and D 0.5 to 10% by weight, yielding lipid-crystalline structures in fluorocarbon (both chloro- and. . . therapeutically active agents, drugs and other materials can be carried into the lungs after release

from and through metered dose **nebulizer**. The spreading agents referred to in the '483 patent are compounds such as the above-described phospholipids, lysophospholipids, plasmalogens, dialklyphospholipids, phosphonolipids, . . .

SUMM . . . glucose, fructose, galactose, pneumogalactan, or dextrose.

Proteins especially suited and advantageously selected for use in the present invention include albumin, pulmonary surfactant specific proteins A or B or C or D, their synthetic analogs, and mixtures thereof.

SUMM . . . in combination: drugs effective in the direct treatment of the subject inflammation such as, for example, corticosteroids including, for example, betamethasone, including, for example, betamethasone dipropionate and betamethasone valerate as well as all other effective formulations; de-congestive agents such as phenylephrine, including, for example, phenylephrine HCL and phenylephrine. . .

SUMM . . . (DPPC/CP) will also produce an effective carrier for this embodiment. If, for example, the therapeutic agent is selected to be betamethasone, the weight ratio of betamethasone to carrier (DPPC/CP) is advantageously selected to be 1 microgram betamethasone to 5 milligrams carrier. However, it has been found that a weight ratio range of 0.5 to 1000 micrograms betamethasone/5 milligrams carrier yields an effective and functional mixture.

DETD . . . were purchased from Sigma Chem., St Louis, Mo. All purchased materials were checked for purity by standard chromatographic analysis. The betamethasone utilized in this example was also purchased from Sigma Chemical. The DPPC and CP were then mixed in the dry powder form in a weight ratio of 200:1 (DPPC:CP). To 5 milligrams of the resultant carrier, 1 microgram of betamethasone was added in order to yield a weight ratio of 5000:1 (carrier: betamethasone). Then 5 grams of this mixture was suspended in 55 grams of the first propellant, trichloromonofluoromethane (P11) and subdivided into. . . The size of the metering valve can be varied to deliver from 1 mg up to 5.4 mg of the DPPC:CP:Betamethasone aerosolized mixture. However, metered dose valves having a greater dosing range are also contemplated and can be utilized in other. .

DETD In the above-described Example "I", wherein the therapeutically active agent is the anti-inflammatory, **betamethasone**, the agent acts directly upon the inflammatory process itself occurring within the upper respiratory epithelium, reducing the production of the. . .

DETD Particle size of the **nebulized** crystals produced and utilized in practicing the present invention is, as discussed below, critical to effective administration. The size (diameter). . . utilizing in a cascade impactor. Flow through the impactor was adjusted to be substantially identical to the flow from a **nebulizer** utilized in practicing the disclosed method. All of the lipid crystals were found to have a diameter equal to or. . .

DETD . . . structure results in, as discussed above, a mean particle size of 1.75 microns. The minute physical dimensions of the individual nebulized particles enables the propellant utilized in practicing the present invention to easily and effectively transfer the disclosed mixture to and. . .

DETD . . . nature of the mixture imparts increased efficiency of particle dispersion within the aerosol mist applied by means of a metered-dose nebulizer. Upon application, the fluorocarbon medium, either chlorofluorocarbon or hydrofluorocarbon, vaporizes rapidly and the DPPC/CP, DPPC/CP drug, DPPC/PG drug or DPPC/PG/CP. . . CLM What is claimed is:

18. The method of claim 1 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.

39. The method of claim 22 wherein the protein is selected from albumin

and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.

44. The method of claim 43 wherein the corticosteroid is **betamethasone** diproprionate, **betamethasone** valerate or combinations thereof.

TT 50-99-7, D-Glucose, biological studies 57-48-7, D-Fructose, biological studies 57-88-5D, Cholesterol, esters 59-23-4, D-Galactose, biological studies 59-42-7, Phenylephrine 61-76-7, Phenylephrine hydrochloride 63-89-8, Dipalmitoylphosphatidylcholine 114-07-8, Erythromycin 303-43-5, Cholesteryl oleate 378-44-9, Betamethasone 601-34-3, Cholesteryl palmitate 2152-44-5, Betamethasone valerate 5593-20-4, Betamethasone dipropionate 17162-39-9, Phenylephrine tartrate 26787-78-0, Amoxicillin 35602-69-8, Cholesteryl stearate 58001-44-8, Clavulanic acid 59277-89-3, Acyclovir 74469-00-4, Augmentin 83905-01-5, Zythromax 534599-12-7, Pneumogalactan (aerosol compns. contg. lipid crystals for decreasing upper respiratory airway resistance)

L12 ANSWER 6 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2003:47500 USPATFULL

TITLE: Composition and method for treatment of otitis externa

INVENTOR(S): Mautone, Alan J., Morristown, NJ, United States

PATENT ASSIGNEE(S): Scientific Development and Research, Inc., Belleville,

NJ, United States (U.S. corporation)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-450884, filed

on 28 Nov 1999, now patented, Pat. No. US 6156294

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Krass, Frederick ASSISTANT EXAMINER: Jagoe, Donna

LEGAL REPRESENTATIVE: Strauss, Esq., Richard L.

NUMBER OF CLAIMS: 123 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT: 1438

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . other phospholipid, and the lysophospholipids; or any of the plasmalogens, dialklylphospholipids, phosphonolipids; carbohydrates and proteins, such as, for example, albumin, **pulmonary**surfactant proteins A, B, C and D. The naturally occurring
surfactant system is further described in U.S. Pat. No. 5,306,483.

SUMM . . . to, glucose, fructose, galactose, pneumogalactan, dextrose, 0.5 to 10% by weight; and proteins such as, but not limited to albumin, pulmonary surfactant specific proteins A, B, C, and D 0.5 to 10% by weight, yielding lipid-crystalline structures in fluorocarbon (both chloro- and. . . therapeutically active agents, drugs and other materials can be carried into the lungs after release from and through metered dose nebulizer. The spreading agents referred to in the '483 patent are compounds such as the above-described phospholipids, lysophospholipids, plasmalogens, dialklyphospholipids, phosphonolipids, . . .

SUMM . . . glucose, fructose, galactose, pneumogalactan, or dextrose. Proteins especially suited and advantageously selected for use in the present invention include albumin, pulmonary surfactant specific proteins A or B or C or D, their synthetic analogs, and mixtures thereof.

SUMM . . . the direct treatment of inflammation such as, for example, corticosteroids including, for example, hydrocortisone, hydrocortisone acetate and dexamethasone sodium phosphate, betamethasone, betamethasone dipropionate and betamethasone valerate as well as all other effective formulations. It is also contemplated that embodiments of the present invention include, as. . .

SUMM . . . (DPPC/CP) will also produce an effective carrier for this

. . . (DPPC/CP) will also produce an effective carrier for this embodiment. If, for example, the therapeutic agent is selected to be betamethasone, the weight ratio of betamethasone to carrier (DPPC/CP) is advantageously selected to be 1 microgram betamethasone to 5 milligrams carrier. However, it has been found that a weight ratio range of 0.5 to 1000 micrograms betamethasone/5 milligrams carrier yields an effective and functional mixture.

DETD Particle size of the **nebulized** crystals produced and utilized in practicing the present invention is, as discussed below, critical to effective administration. The size (diameter). . . utilizing in a cascade impactor. Flow through the impactor was adjusted to be substantially identical to the flow from a **nebulizer** utilized in practicing the disclosed method. All of the lipid crystals were found to have a diameter equal to or. . .

DETD . . . structure results in, as discussed above, a mean particle size of 1.75 microns. The minute physical dimensions of the individual nebulized particles enables the propellant utilized in practicing the present invention to easily and effectively transfer the disclosed mixture to and. . .

DETD . . . nature of the mixture imparts increased efficiency of particle dispersion within the aerosol mist applied by means of a metered-dose nebulizer. Upon application, the fluorocarbon medium, either chlorofluorocarbon or hydrofluorocarbon, vaporizes rapidly and the DPPC/CP, DPPC/CP drug, DPPC/PG drug or DPPC/PG/CP. . . CLM What is claimed is:

17. The method of claim 1 wherein the protein is selected from the group consisting of albumin and **pulmonary surfactant** specific proteins A, B, C, D and mixtures thereof.

- 37. The method of claim 21 wherein the protein is selected from the group consisting of albumin and **pulmonary surfactant** specific proteins A, B, C, D and mixtures thereof.
- . method of claim 41 wherein the corticosteroid is selected from the group consisting of hydrocortisone, hydrocortisone acetate, dexamethasone sodium phosphate, betamethasone, betamethasone diproprionate, betamethasone valerate and combinations thereof.
 - 64. The process of claim 49 wherein the protein is selected from the group consisting of albumin and **pulmonary surfactant** specific proteins A, B, C, D and mixtures thereof.
 - 85. The process of claim 70 wherein the protein is selected from the group consisting of albumin and **pulmonary surfactant** specific proteins A, B, C, D and mixtures thereof.
- . process of claim 94 wherein the corticosteroid is selected from the group consisting of hydrocortisone, hydrocortisone acetate, dexamethasone sodium phosphate, betamethasone, betamethasone dipropionate, betamethasone valerate and combinations thereof.
 - 119. The method of claim 103 wherein the protein is selected from the group consisting of albumin and **pulmonary surfactant** specific proteins A, B, C, D and mixtures thereof.

50-02-2, Dexamethasone 50-03-3, Hydrocortisone acetate 50-23-7, IT Hydrocortisone 378-44-9, Betamethasone 1264-72-8, Colistin 1400-61-9, Nystatin 1404-26-8, Polymyxin b 1405-10-3, Neomycin sulfate 2152-44-5, Betamethasone valerate 5593-20-4, 23593-75-1, 7632-05-5, Sodium phosphate Betamethasone dipropionate Clotrimazole 59277-89-3, Acyclovir (treatment of otitis externa with aerosol formulation contg. medicaments such as antibiotics, corticosteroids, antivirals, and nucleic acids)

L12 ANSWER 7 OF 11 USPATFULL on STN

ACCESSION NUMBER:

2002:171597 USPATFULL

TITLE:

Composition and method for decreasing upper respiratory

airway resistance

INVENTOR(S):

Mautone, Alan J., Morristown, NJ, UNITED STATES

| | NUMBER | KIND | DATE | |
|---------------------|---------------|------|----------|--|
| | | | | |
| PATENT INFORMATION: | US 2002090344 | A1 | 20020711 | |
| APPLICATION INFO.: | US 2001-11994 | A1 | 20011204 | |

(10)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2000-639739, filed on 16 Aug 2000, PENDING Continuation-in-part of Ser.

No. US 1999-450884, filed on 28 Nov 1999, PATENTED

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE:

Richard L. Strauss, Esq., 2492 Oceanside Road,

Oceanside, NY, 11572

NUMBER OF CLAIMS: 133 EXEMPLARY CLAIM: 1740 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

. . other phospholipid, and the lysophospholipids; or any of the SUMM plasmalogens, dialklylphospholipids, phosphonolipids; carbohydrates and proteins, such as, for example, albumin, pulmonary surfactant proteins A, B, C and D. The naturally occurring surfactant system is further described in U.S. Pat. No. 5,306,483.

SUMM . to, glucose, fructose, galactose, pneumogalactan, dextrose, 0.5 to 10% by weight; and proteins such as, but not limited to albumin, pulmonary surfactant specific proteins A, B, C, and D 0.5 to 10% by weight, yielding lipid-crystalline structures in fluorocarbon (both chloro- and. . . active agents, drugs and other materials can be carried into the lungs after release from and through a metered dose nebulizer. The spreading agents referred to in the '483 patent are compounds such as the above-described phospholipids, lysophospholipids, plasmalogens, dialklyphospholipids, phosphonolipids,.

glucose, fructose, galactose, pneumogalactan, or dextrose. SUMM Proteins especially suited and advantageously selected for use in the present invention include albumin, pulmonary surfactant specific proteins A or B or C or D, their synthetic analogs, and mixtures thereof.

. in combination: drugs effective in the direct treatment of the SUMM subject inflammation such as, for example, corticosteroids including, for example, betamethasone, including, for example, betamethasone dipropionate and betamethasone valerate as well as all other effective formulations; de-congestive agents such as phenylephrine, including, for example, phenylephrine HCL and phenylephrine.

SUMM . (DPPC/CP) will also produce an effective carrier for this embodiment. If, for example, the therapeutic agent is selected to be betamethasone, the weight ratio of betamethasone to carrier (DPPC/CP) is advantageously selected to be 1 microgram betamethasone to 5 milligrams carrier. However, it has been found that a weight ratio range of 0.5 to 1000 micrograms

betamethasone/5 milligrams carrier yields an effective and functional mixture.

. were purchased from Sigma Chem., St Louis, Mo. All purchased DETD materials were checked for purity by standard chromatographic analysis. The betamethasone utilized in this example was also purchased from Sigma Chemical. The DPPC and CP were then mixed in the dry powder form in a weight ratio of 200:1 (DPPC:CP). To 5 milligrams of the resultant carrier, 1 microgram of betamethasone was added in order to yield a weight ratio of 5000:1 (carrier: betamethasone). Then 5 grams of this mixture was suspended in 55 grams of the first propellant, trichloromonofluoromethane (P11) and subdivided into. . The size of the metering valve can be varied to deliver from 1 mg up to 5.4 mg of the DPPC:CP:Betamethasone aerosolized mixture. However, metered dose valves having a greater dosing range are also contemplated and can be utilized in other.

- DETD [0070] In the above-described Example "I", wherein the therapeutically active agent is the anti-inflammatory, betamethasone, the agent acts directly upon the inflammatory process itself occurring within the upper respiratory epithelium, reducing the production of the.
- [0073] Particle size of the nebulized crystals produced and DETD utilized in practicing the present invention is, as discussed below, important for effective administration. The size (diameter). utilizing in a cascade impactor. Flow through the impactor was adjusted to be substantially identical to the flow from a nebulizer utilized in practicing the disclosed method. All of the lipid crystals were found to have a diameter equal to or.
- DETD structure results in, as discussed above, a mean particle size of 1.75 microns. The minute physical dimensions of the individual nebulized particles enables the propellant utilized in practicing the present invention to easily and effectively transfer the disclosed mixture to and.
- . nature of the mixture imparts increased efficiency of particle DETD dispersion within the aerosol mist applied by means of a metered-dose nebulizer. For example, upon application, the fluorocarbon medium, either chlorofluorocarbon or hydrofluorocarbon, vaporizes rapidly and the DPPC/CP, DPPC/CP drug, DPPC/PG drug. CLMWhat is claimed is:
 - 14. The method of claim 1 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 34. The method of claim 21 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 37. The method of claim 36 wherein said anti-inflammatory agent is betamethasone.
 - 57. The process of claim 45 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 76. The process of claim 64 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 79. The process of claim 78 wherein the anti-inflammatory agent is selected to be betamethasone.
 - 100. The method of claim 87 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.

- 103. The method of claim 102 wherein said anti-inflammatory agent is betamethasone.
- 123. The process of claim 111 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
- 126. The process of claim 125 wherein said anti-inflammatory agent is betamethasone.
- 50-99-7, D-Glucose, biological studies 57-10-3, Palmitic acid, IT biological studies 57-48-7, Fructose, biological studies 57-87-4, 57-88-5, Cholesterol, biological studies 57-88-5D, Cholesterol, esters 59-23-4, Galactose, biological studies 59-42-7, 67-97-0, Cholecalciferol 112-80-1, Oleic acid, Phenylephrine biological studies 114-07-8, Erythromycin 303-43-5, Cholesteryl oleate 378-44-9, Betamethasone 601-34-3, Cholesteryl palmitate 2644-64-6, 1,2 Dipalmitoylphosphatidylcholine 26787-78-0, Amoxicillin 35602-69-8, Cholesteryl stearate 74469-00-4, Augmentin 83905-01-5

(compn. and method for decreasing upper respiratory airway resistance using aerosolized lipid crystals)

L12 ANSWER 8 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2002:148246 USPATFULL

TITLE: Composition and method for treatment of otitis external

INVENTOR(S): Mautone, Alan J., Morristown, NJ, UNITED STATES

| | NUMBER | KIND | DATE | |
|---------------------|---------------|------|----------|-----|
| PATENT INFORMATION: | US 2002076383 | A1 | 20020620 | |
| APPLICATION INFO.: | US 2001-11626 | A1 | 20011211 | (|
| | | | O 37 - | TTC |

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 2000-639730, filed on 16 Aug 2000, PENDING Continuation-in-part of Ser.

No. US 1999-450884, filed on 28 Nov 1999, PATENTED

(10)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Richard L. Strauss, Esq., 2492 Oceanside Road,

Oceanside, NY, 11572

NUMBER OF CLAIMS: 130 EXEMPLARY CLAIM: 1 LINE COUNT: 1640

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . other phospholipid, and the lysophospholipids; or any of the plasmalogens, dialklylphospholipids, phosphonolipids; carbohydrates and proteins, such as, for example, albumin, pulmonary surfactant proteins A, B, C and D. The naturally occurring surfactant system is further described in U.S. Pat. No. 5,306,483.

SUMM . . . to, glucose, fructose, galactose, pneumogalactan, dextrose, 0.5 to 10% by weight; and proteins such as, but not limited to albumin, pulmonary surfactant specific proteins A, B, C, and D 0.5 to 10% by weight, yielding lipid-crystalline structures in fluorocarbon (both chloro- and. . . therapeutically active agents, drugs and other materials can be carried into the lungs after release from and through metered dose nebulizer. The spreading agents referred to in the '483 patent are compounds such as the above-described phospholipids, lysophospholipids, plasmalogens, dialklyphospholipids, phosphonolipids, . . .

SUMM . . . glucose, fructose, galactose, pneumogalactan, or dextrose.

Proteins especially suited and advantageously selected for use in the present invention include albumin, pulmonary surfactant specific proteins A or B or C or D, their synthetic analogs, and mixtures thereof.

DETD . . . the direct treatment of inflammation such as, for example,

embodiment. If, for example, the therapeutic agent is selected to be betamethasone, the weight ratio of betamethasone to carrier (DPPC/CP) is advantageously selected to be 1 microgram betamethasone to 5 milligrams carrier. However, it has been found that a weight ratio range of 0.5 to 1000 micrograms betamethasone/5 milligrams carrier yields an effective and functional mixture.

DETD

DETD . . . utilizing in a cascade impactor. Flow through the impactor was adjusted to be substantially identical to the flow from a nebulizer utilized in practicing the disclosed method. All of the lipid crystals were found to have a diameter equal to or. . .

DETD . . . structure results in, as discussed above, a mean particle size of 1.75 microns. The minute physical dimensions of the individual nebulized particles enables the propellant utilized in practicing the present invention to easily and effectively transfer the disclosed mixture to and. . .

DETD . . . nature of the mixture imparts increased efficiency of particle dispersion within the aerosol mist applied by means of a metered-dose nebulizer. Upon application, the fluorocarbon medium, either chlorofluorocarbon or hydrofluorocarbon, vaporizes rapidly and the DPPC/CP, DPPC/CP drug, DPPC/PG drug or DPPC/PG/CP. . . CLM What is claimed is:

13. The method of claim 1 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.

- 32. The method of claim 20 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
- 35. The method of claim 34 wherein said anti-inflammatory agent is betamethasone.
- 55. The process of claim 43 wherein the protein is selected from albumin and **pulmonary surfactant** specific proteins A or B or C or D or mixtures thereof.
- 74. The process of claim 62 wherein the protein is selected from albumin and **pulmonary surfactant** specific proteins A or B or C or D or mixtures thereof.
- 77. The process of claim 76 wherein the anti-inflammatory agent is selected to be **betamethasone**.
- 97. The method of claim 85 wherein the protein is selected from albumin and **pulmonary surfactant** specific proteins A or B or C or D or mixtures thereof.
- 100. The method of claim 99 wherein said anti-inflammatory agent is betamethasone.
- 120. The process of claim 108 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
- 123. The process of claim 122 wherein said anti-inflammatory agent is betamethasone.

L12 ANSWER 9 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2002:125993 USPATFULL

TITLE: Composition and method for treatment of otitis media INVENTOR(S): Mautone, Alan J., Morristown, NJ, UNITED STATES

PATENT INFORMATION: US 2002064503 A1 20020530 APPLICATION INFO.: US 2001-11344 A1 20011204 (10)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 2000-639682, filed

on 16 Aug 2000, PENDING Continuation of Ser. No. US

1999-450884, filed on 28 Nov 1999, PATENTED

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Richard L. Strauss, Esq., 2492 Oceanside Road,

Oceanside, NY, 11572

NUMBER OF CLAIMS: 133
EXEMPLARY CLAIM: 1
LINE COUNT: 1671

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . phospholipid, and of the lysophospholipids; or any of the plasmalogens, dialklylphospholipids, phosphonolipids, carbohydrates and proteins, such as, for example, albumin, pulmonary surfactant proteins A, B, C and D. The naturally occurring

surfactant system is further described in U.S. Pat. No. 5,306,483.

SUMM . . . galactose, pneumogalactan, dextrose (or mixtures thereof), 0.5 to 10% by weight, and proteins such as, but not limited to albumin, pulmonary surfactant specific proteins A, B, C, and D 0.5 to 10% by weight, compounds in lipid-crystalline structures in fluorocarbon (both chloro-. . . therapeutically active agents, drugs and other materials can be carried into the lungs after release from and through metered dose nebulizer. The spreading agents referred to in the '483 patent are compounds such as the above-described phospholipids, lysophospholipids, plasmalogens, dialklyphospholipids, phosphonolipids, . .

SUMM . . . galactose, pneumogalactan, dextrose or mixtures thereof.

Proteins especially suited and advantageously selected for use in the present invention include albumin, pulmonary surfactant specific proteins A or B or C or D, their synthetic analogs, and mixtures thereof.

DETD . . . refers to those drugs effective in treatment of otitis media including, but not limited to anti-inflammatory agents including, for example, betamethasone, including, for example, betamethasone dipropionate and betamethasone valerate as well as all other effective formulations; de-congestive agents such as phenylephrine, including, for example, phenylephrine HCL and phenylephrine. . .

DETD . . . (DPPC/CP) will also produce an effective carrier for this embodiment. If, for example, the therapeutic agent is selected to be betamethasone, the weight ratio of betamethasone to carrier (DPPC/CP) is advantageously selected to be 1 microgram betamethasone to 5 milligrams carrier. However, it has been found that a weight ratio range of 0.5 to 1000 micrograms betamethasone/5 milligrams carrier yields an effective and functional mixture.

DETD . . . were purchased from Sigma Chem., St Louis, Mo. All purchased materials were checked for purity by standard chromatographic analysis. The betamethasone utilized in this example was also purchased from Sigma Chemical. The DPPC and CP were then mixed in the dry powder form in a weight ratio of 200:1 (DPPC:CP). To 5 milligrams of the resultant carrier, 1 microgram of betamethasone was added in order to yield a weight ratio of 5000:1 (carrier: betamethasone). Then 5 grams of this mixture was suspended in 55 grams of the first

- propellant, trichloromonofluoromethane (P11) and subdivided into. . . DETD . . . The size of the metering valve can be varied to deliver from 1 mg up to 5.4 mg of the DPPC:CP:Betamethasone aerosolized mixture. However, metered dose valves having a greater dosing range are also contemplated and can be utilized in other. . .
- DETD [0066] In the above-described Example "I", wherein the therapeutically active agent is the anti-inflammatory betamethasone, the agent acts directly upon the auditory tube itself, reducing the excess mucoid secretions and swelling of the auditory tube. . .
- DETD . . . in an aerosolized metered dose inhaler (MDI) viz 1) Placebo (normal saline); 2) Surfactant alone (DPPC:CP (200:1); 3) Surfactant with betamethasone (5 mg carrier to 10 micrograms betamethasone diproprionate); 4) Surfactant with phenylephrine (995 mg carrier to 160 micrograms phenylephrine HCI). In-vivo Typanometry and Micro-otoscopy was done on. . . after the development of OME. Resolution of OME was observed by micro-otoscopy on the 6.sup.th, day in the surfactant with betamethasone group, on the 10.sup.th day with the surfactant alone group, and on the 16.sup.th day for all other groups. The. . .
- DETD [0077] Particle size of the **nebulized** crystals produced and utilized in practicing the present invention is, as discussed below, is important for effective administration. The size. . . determined utilizing a cascade impactor. Flow through the impactor was adjusted to be substantially identical to the flow from a **nebulizer** utilized in practicing the disclosed method. All of the lipid crystals were found to have a diameter equal to or. . .
- DETD . . . structure results in, as discussed above, a mean particle size of 1.75 microns. The minute physical dimensions of the individual nebulized particles enables the propellant utilized in practicing the present invention to easily and effectively transfer the disclosed mixture to and. . .
- DETD . . . nature of the mixture imparts increased efficiency of particle dispersion within the aerosol mist applied by means of a metered-dose nebulizer. Upon application, the propellant, such as, for example a fluorocarbon medium, (either chlorofluorocarbon or hydrofluorocarbon), vaporizes rapidly and the DPPC/CP, . .
- CLM What is claimed is:
 14. The method of claim 1. wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 34. The method of claim 21 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 37. The method of claim 36 wherein said anti-inflammatory agent is betamethasone.
 - 57. The process of claim 45 wherein the protein is selected from albumin and **pulmonary surfactant** specific proteins A or B or C or D or mixtures thereof.
 - 76. The process of claim 64 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 79. The process of claim 78 wherein the anti-inflammatory agent is selected to be **betamethasone**.
 - 100. The method of claim 87 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 103. The method of claim 102 wherein said anti-inflammatory agent is

betamethasone.

123. The process of claim 111 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.

126. The process of claim 125 wherein said anti-inflammatory agent is betamethasone.

L12 ANSWER 10 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2000:164054 USPATFULL

TITLE: Composition and method for treatment of otitis media

INVENTOR(S): Mautone, Alan J., Morristown, NJ, United States

PATENT ASSIGNEE(S): Scientific Development and Research, Inc., Belleville,

NJ, United States (U.S. corporation)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Krass, Frederick ASSISTANT EXAMINER: Jagoe, Donna

LEGAL REPRESENTATIVE: Strauss, Esq., Richard L.

NUMBER OF CLAIMS: 46
EXEMPLARY CLAIM: 1
LINE COUNT: 1128

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . well as any of the lysophospholipids; any of the plasmalogens, dialklylphospholipids, phosphonolipids, carbohydrates; and proteins, such as, for example, albumin, pulmonary surfactant proteins A, B, C and D. The naturally occurring surfactant system is further described in U.S. Pat. No. 5,306,483.

SUMM . . . to, glucose, fructose, galactose, pneumogalactan, dextrose, 0.5 to 10% by weight, and proteins such as, but not limited to albumin, pulmonary surfactant specific proteins A, B, C, and D 0.5 to 10% by weight, compounds in lipid-crystalline structures in fluorocarbon (both chloro- . . therapeutically active agents, drugs and other materials can be carried into the lungs after release from and through metered dose nebulizer. The spreading agents referred to in the '483 patent are compounds such as the above-described phospholipids, lysophospholipids, plasmalogens, dialklyphospholipids, phosphonolipids, . .

SUMM . . . glucose, fructose, galactose, pneumogalactan, or dextrose. Proteins especially suited and advantageously selected for use in the present invention include albumin, pulmonary surfactant specific proteins A or B or C or D, their synthetic analogs, and mixtures thereof.

SUMM . . . refers to those drugs effective in treatment of otitis media including, but not limited to anti-inflammatory agents including, for example, betamethasone, including, for example, betamethasone dipropionate and betamethasone valerate as well as all other effective formulations; de-congestive agents such as phenylephrine, including, for example, phenylephrine HCL and phenylephrine. . .

SUMM . . . (DPPC/CP) will also produce an effective carrier for this embodiment. If, for example, the therapeutic agent is selected to be

betamethasone, the weight ratio of betamethasone to carrier (DPPC/CP) is advantageously selected to be 1 microgram betamethasone to 5 milligrams carrier. However, it has been found that a weight ratio range of 0.5 to 1000 micrograms betamethasone/5 milligrams carrier yields an effective and functional mixture.

DETD . . . were purchased from Sigma Chem., St Louis, Mo. All purchased materials were checked for purity by standard chromatographic analysis. The betamethasone utilized in this example was also purchased from Sigma Chemical. The DPPC and CP were then mixed in the dry powder form in a weight ratio of 200:1 (DPPC:CP). To 5 milligrams of the resultant carrier, 1 microgram of betamethasone was added in order to yield a weight ratio of 5000:1 (carrier: betamethasone). Then 5 grams of this mixture was suspended in 55 grams of the first propellant, trichloromonofluoromethane (P11) and subdivided into. . . The size of the metering valve can be varied to deliver from 1 mg up to 5.4 mg of the DPPC:CP:Betamethasone aerosolized mixture. However, metered dose valves having a greater dosing range are also contemplated and can be utilized in other. . .

- DETD In the above-described Example "I", wherein the therapeutically active agent is the anti-inflammatory, **betamethasone**, the agent acts directly upon the auditory tube itself, reducing the excess mucoid secretions and swelling of the auditory tube. . .
- DETD . . . in an aerosolized metered dose inhaler (MDI) viz 1) Placebo (normal saline); 2) Surfactant alone (DPPC:CP (200:1); 3) Surfactant with betamethasone (5 mg carrier to 10 micrograms betamethasone diproprionate); 4) Surfactant with phenylephrine (995 mg carrier to 160 micrograms phenylephrine HCl). In-vivo Typanometry and Micro-otoscopy was done on. . . after the development of OME. Resolution of OME was observed by micro-otoscopy on the 6.sup.th, day in the surfactant with betamethasone group, on the 10.sup.th day with the surfactant alone group, and on the 16.sup.th day for all other groups. The. . .
- DETD Particle size of the **nebulized** crystals produced and utilized in practicing the present invention is, as discussed below, critical to effective administration. The size (diameter). . . utilizing in a cascade impactor. Flow through the impactor was adjusted to be substantially identical to the flow from a **nebulizer** utilized in practicing the disclosed method. All of the lipid crystals were found to have a diameter equal to or. . .
- DETD . . . structure results in, as discussed above, a mean particle size of 1.75 microns. The minute physical dimensions of the individual nebulized particles enables the propellant utilized in practicing the present invention to easily and effectively transfer the disclosed mixture to and. . .
- DETD . . . nature of the mixture imparts increased efficiency of particle dispersion within the aerosol mist applied by means of a metered-dose nebulizer. Upon application, the fluorocarbon medium, either chlorofluorocarbon or hydrofluorocarbon, vaporizes rapidly and the DPPC/CP, DPPC/CP drug, DPPC/PG drug or DPPC/PG/CP. . .
- CLM What is claimed is:
 17. The method of claim 1 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 37. The method of claim 21 wherein the protein is selected from albumin and pulmonary surfactant specific proteins A or B or C or D or mixtures thereof.
 - 41. The method of claim 40 wherein said anti-inflammatory agent is betamethasone.
- IT 114-07-8, Erythromycin **378-44-9**, Betamethasone 26787-78-0, Amoxicillin 74469-00-4, Augmentin 83905-01-5, Zithromax

(lipid aerosols for treatment of otitis media)

L12 ANSWER 11 OF 11 USPATFULL on STN ACCESSION NUMBER: 96:57206 USPATFULL Use of liquid fluorocarbons to facilitate pulmonary TITLE: drug delivery Rosenberg, Gwen H., Rancho Santa Fe, CA, United States INVENTOR(S): PATENT ASSIGNEE(S): Alliance Pharmaceutical Corp., San Diego, CA, United States (U.S. corporation) NUMBER KIND DATE _____ PATENT INFORMATION: US 5531219 19960702 US 1994-334688 19941104 (8) APPLICATION INFO.: DOCUMENT TYPE: Utility Granted FILE SEGMENT: Lewis, Aaron J. PRIMARY EXAMINER: LEGAL REPRESENTATIVE: Knobbe, Martens, Olson & Bear NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: 1248 LINE COUNT: CAS INDEXING IS AVAILABLE FOR THIS PATENT. . . States accounting for up to 5,000 infant deaths annually. The primary etiology of RDS is attributed to insufficient amounts of pulmonary surfactant. Premature infants born before the 36th week of gestation are at greatest risk because of insufficient lung development. Neonates born. . . . powdered form, in microcrystalline suspension, in a clathrate SUMM with other compounds, in an aerosol, in a gaseous phase, in a nebulized suspension or any other form of small particles that can be suspended in a gas that is well known in. . . anti-inflammatory agents including triamcinolone DETD (9-fluoro-11.beta., 16.alpha., 17,21-tetrahydroxypregna-1,4-diene-3,20dione), triamcinolone acetonide (9-fluoro-11.beta., 16.alpha., 17,21-tetrahydroxypregna-1,4-diene-3,20-dione cyclic 16, 17-acetal), beclomethasone dipropionate (9-chloro-11.beta., 17,21-trihydroxy-16.beta.-methylpregna-1,4-diene-3,20-dione 17,21-dipropionate), betamethasone sodium phosphate (9-fluoro-11.beta., 17,21-trihydroxy-16.beta.-methylpregna-1,4-diene-3,20-dione 21-sodium

phosphate), hydrocortisone (pregna-4-ene-3,20-dione, 21 (acetyloxy)-11,

oxy)pregna-1,4-diene-3,20-dione 17,21-disodium salt), and triamcinolone.

(9-fluoro-11.beta., 17-dihydroxy-16.alpha.-methyl-21-(phosphono-

17-dihydroxy-acetate), dexamethasone sodium phosphate